CLAIMS

1. A compound of Formula (I):

5 wherein:

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One of R1 and R2 is H and the other represents – NHCONHR4

wherein R^4 represents a phenyl or naphthyl group (which may be optionally substituted by one or more substituents independently selected from -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, - CH₂CH₂CH₂-, halogen, C₁₋₆ alkoxy, C₁₋₆ haloalkoxy, OH, NO₂). C₃₋₇ cycloalkyl or R^4 together with the NH to which it is bonded forms a morpholino group and

 R^3 is H or NHR⁵ wherein R^5 is H, -quinolinyl or -isoquinolinyl, -(CONH)_p phenyl (wherein p is 0 or 1 and the phenyl is optionally substituted by one or more substituents independently selected from halogen, -C₁₋₈ alkyl, -C₁₋₈ haloalkyl, -morpholino, -SO₂NH₂, benzothiazole (substituted by methyl))

or a salt, solvate, or physiologically functional derivative thereof.

- 2. A compound according to claim 1 wherein R⁴ represents a phenyl group (which may be optionally substituted by one or more substituents selected from -C₁₋₈ haloalkyl, -CH₂CH₂CH₂-, halogen) or C₃₋₇ cycloalkyl.
- A compound according to claims 1 2 wherein R³ is H or –NH R⁵ where in R⁵ is H, quinolinyl, -(CONH)p phenyl (wherein p is 0 or 1 and the phenyl is optionally substituted by one or more substituents independently selected from halogen, -C₁₋₆ haloalkyl –morpholino, -SO₂NH₂, benzothiazole, (substituted by methyl)).

4. A compound according to claims 1 – 3 of formula (1a)

$$\mathbb{R}^{8}$$
 (1a)

wherein one of R⁶ and R⁷ is H and the other represents -NHCONHR⁹;

R⁹ represents a phenyl group (which may be optionally substituted by one or more substituents independently selected from -C₁₋₆ haloalkyl, -CH₂CH₂-, halogen) or C₃₋₇ cycloalkyl;

R⁸ is H or NHR¹⁰;

R¹⁰ is H quinolinyl, -(CONH)p phenyl (where p is 0 or 1 and the phenyl is optionally substituted by one or more substituents independently selected from halogen, -C₁₋₆ haloalkyl, -morpholino, -SO₂NH₂, benzothiazole (substituted by methyl)).

5. A compound according to claim 4 wherein NHCONHR⁹ represents

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6. A compound according to claim 4 and 5 where in R¹⁰ is H,

- 7. A compound as claimed in claim 1 6, selected from the group consisting of: 1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-(3-isoquinolin-5-ylphenyl)urea;
- 5 1-Cyclohexyl-3-(3-isoquinolin-5-ylphenyl)urea;
 - 1-[3-(1-Amino-isoquinolin-5-yl)-phenyl]-3-(2-fluoro-5-trifluoromethyl-phenyl)-urea;
 - 1-(2-fluoro-5-trifluoromethyl-phenyl)-3-(5-{3-[3-(2-fluoro-5-trifluoromethyl-phenyl)-ureidol-phenyl}-isoquinolin-1-yl)-urea;
 - 1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-{3-[1-(quinolin-6-ylamino)-isoquinolin-5-yl]-
- 10 phenyi}-urea;

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- 1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-(4-{1-[4-(6-methyl-benzothiazol-2-yl)-phenylamino]-isoquinolin-5-yl}-phenyl)-urea;
- 1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-(3-{1-[4-(6-methyl-benzothiazol-2-yl)-phenylamino]-isoquinolin-5-yl}-phenyl)-urea;
- 15 1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-(4-isoquinolin-5-ylphenyl)urea;
 - 1-Indan-5-yl-3-(3-isoquinolin-5-yl-phenyl)-urea;
 - 1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-{3-[1-(4-morpholin-4-yl-phenylamino)-isoquinolin-5-yl]-phenyl}-urea;
 - 3-{5-[3-(3-Cyclohexyl-ureido)-phenyl]-isoquinolin-1-ylamino}-benzenesulfonamide;

or a salt, solvate, or physiologically functional derivative thereof.

8. A pharmaceutical composition, comprising: a therapeutically effective amount of a compound as claimed in any one of claims 1 - 7, or a salt, solvate, or a physiologically functional derivative thereof and one or more of pharmaceutically acceptable carriers, diluents and excipients.

- 9. A pharmaceutical composition according to claim 8 further comprising an agent to inhibit growth factor receptor function
- 10. A compound as claimed in any of claims 1 7, or a salt, solvate, or a physiologically functional derivative thereof for use in therapy.
 - 11. A method of treating a disorder in a mammal, said disorder being mediated by at least one of inappropriate TIE-2, Eph B4 and VEGFR-2 activity, comprising administering to said mammal a compound according to claims 1 7 or a salt, solvate or a physiologically functional derivative thereof.

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- 12. The use of a compound according to claims 1 7, or a salt, solvate, or a physiologically functional derivative thereof in the manufacture of a medicament for use in the treatment of a disorder mediated by at least one of inappropriate TIE-2, EphB4 and VEGFR-2 activity.
- 13. A method of treating a disorder in a mammal, said disorder being mediated by at least one of inappropriate TIE-2, Eph B4 and VEGFR-2 activity, comprising: administering to said mammal (i) a compound according to claims 1 7, or a salt, solvate or physiologically functional derivative thereof and (ii) an agent to inhibit growth factor receptor function.
- 14. The use of a compound according to claims 1 7, or a salt, solvate or physiologically functional derivative thereof and an agent to inhibit growth factor receptor function in the manufacture of a medicament for the treatment of a disorder mediated by at least one of inappropriate TIE-2, EphB4 and VEGFR2 activity.
- 15. A method of treating a disorder in a mammal, said disorder beingcharacterized by inappropriate angiogenesis, comprising administering to said mammal a compound according to claims 1 7, or a salt, solvate or physiologically functional derivative thereof.
- 16. The use of a compound according to claims 1 7 or a salt, solvate or physiologically functional derivative thereof in the manufacture of a medicament for the treatment of inappropriate angiogenesis.